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Bidirectional Transfer Study of Polystyrene Nanoparticles across the Placental Barrier in an *ex Vivo* Human Placental Perfusion Model

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Materials and Methods

Cell culture

MTS viability assay

Ex vivo human placental perfusion model

Antipyrine transfer

Viability and functionality of the placenta

Histopathological evaluation

References

Figure S1. Stability (A) and leakage (B) of the fluorescence dye was assessed after 3, 6, 24, 48 and 72 hrs of incubation at 37°C in perfusion medium. Shown are the mean percentages of initial (time point 0) fluorescence intensity \pm SD (A) and the free fluorescence of the large PS particles as percentage of total fluorescence intensity \pm SD (B), n=3. (C) Representative TEM micrographs of plain 50 nm, 240 nm and carboxylate-modified 300 nm PS beads in the fetal or maternal circulation after 1.5 - 6 hrs of normal (M to F direction) or reverse (F to M direction) perfusion.

Figure S2. BeWo cell viability after 3 (A) and 24 hrs (B) of exposure to PS beads measured with a MTS assay. CdSO₄ served as positive control for cytotoxicity. Data represent the mean percentages of viable cells compared to the untreated control \pm SD of at least 3 independent experiments.

Figure S3. (A + B) Fetal-to-maternal (F/M) or maternal-to-fetal transfer ratio (M/F) of radiolabeled ¹⁴C-antipyrine over a time period of 6 hrs of normal (A) or reverse (B) placenta perfusion with or without PS beads. (C + D) Glucose consumption and lactate production (C) and human chorionic gonadotropin (hCG) and leptin hormone production rate (D) during *ex vivo* human placental perfusion with or without particles (control). All data is expressed as the mean \pm SD of at least 3 independent experiments.